WEST Search History

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DATE: Tuesday, February 20, 2007

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	DB=PGI	PB, USPT, USOC, EPAB, JPAB, DWPI; PLUR=YES	S; OP=ADJ
	L6	L1 same (mutant? or variant? or mutation?)	4
	L5	homoserine transsuccinylase mutant?	1
	L4	L1 and (mutant? or variant? or mutation?)	16
	L3	homoserine transsuccinylase.clm.	1
	L2	homoserine transsuccinylase	23
	L1	homoserine transsuccinylase	23

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WEST Search History

DATE: Tuesday, February 20, 2007

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	DB=PGP	B, USPT, USOC, EPAB, JPAB, DWPI; PLUR=YES	S; OP=ADJ
	L9	L8 and (mutation? or mutant? or variant?)	6
	L8	homoserine succinyltransferase	6
	L7	homoserine succinyltransferase mutant?	0
	DB=PGP	B; PLUR=YES; OP=ADJ	
	L6	L4 and metA	1
	L5	L4 and homoserine transsuccinylase	0
	L4	20020106800	1
	DB = USP'	T; PLUR=YES; OP=ADJ	
	L3	L1 and homoserine transsuccinylase	0
	L2	L1 and metA	0
<u>`</u>	L1	5120837	7

END OF SEARCH HISTORY

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L3: Entry 1 of 1

File: PGPB

Jul 20, 2006

PGPUB-DOCUMENT-NUMBER: 20060160173

PGPUB-FILING-TYPE:

DOCUMENT-IDENTIFIER: US 20060160173 A1

TITLE: Feedback-resistant homoserine transsuccinylases having a modified c terminus

PUBLICATION-DATE: July 20, 2006

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY
Leonhartsberger; Susanne	Munchen		DE .
Winterhalter; Christoph	Pocking		DE
Pfeiffer; Kerstin	Munchen		DE
Bauer; Brigitte	Munchen		DE

US-CL-CURRENT: 435/69.1; 435/193, 435/252.3, 435/471, 536/23.2

CLAIMS:

- 1. A homoserine transsuccinylase which, as compared with a homoserine transsuccinylase wild-type enzyme, exhibits a reduced sensitivity toward L-methionine or SAM, with the wild-type enzyme possessing an amino acid sequence which comprises a constituent sequence TyrGlnXaaThrPro, with the Thr of this constituent sequence being between position 285 and 310 of the amino acid sequence and with position 1 being the starting methionine, wherein it exhibits a change of at least 2 amino acids as compared with the wild-type enzyme, with this change being in the Thr of the constituent sequence or C-terminally thereof.
- 2. A <u>homoserine transsuccinylase</u> as claimed in claim 1, wherein it exhibits a change of at least 5 amino acids, preferably of at least 10 amino acids.
- 3. A <u>homoserine transsuccinylase</u> as claimed in claim 1, wherein it exhibits a resistance toward the inhibitors SAM and/or L-methionine which is increased (increased Ki) at least 2-fold as compared with that of the wild-type enzyme.
- 4. A <u>homoserine transsuccinylase</u> as claimed in claim 1, wherein it contains one of the mutations listed in Table 1.
- 5. A metA allele which encodes a homoserine transsuccinylase as claimed in claim 1.
- 6. A plasmid, wherein it contains a metA allele as claimed in claim 5 together with a promoter.
- 7. A microorganism strain, wherein it contains a feedback-resistant metA allele as claimed in claim 5.
- 8. A microorganism strain as claimed in claim 7, wherein it is a Gram-negative bacterial strain, preferably E. coli.

 $\stackrel{\searrow}{\mathcal{N}}$ A method for preparing L-methionine or SAM by culturing a microorganism strain as claimed in claim 7.

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=> file medline hcaplus biosis embase biotechds scisearch
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L2 9 DUP REM L1 (5 DUPLICATES REMOVED)

=> s 12 and (?101? or ? 294?)
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=> s 12 and (101? or 294?)
TERM '294?' EXCEEDED TRUNCATION LIMITS - SEARCH ENDED
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=> s 12 and 294

L5 0 L2 AND 294

=> d 12 1-9 ibib ab

L2 ANSWER 1 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2007:88482 HCAPLUS

TITLE:

Use of dimethyl disulfide for methionine production in microorganisms

INVENTOR(S): Zelder, Oskar; Haefner, Stefan; Herold, Andrea;

Klopprogge, Corinna; Schroder, Hartwig; Yocum, R.

Rogers; Williams, Mark K.

PATENT ASSIGNEE(S):

SOURCE:

BASF A.-G., Germany PCT Int. Appl., 99pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PAT	PATENT NO.				KIND DATE			APPLICATION NO.					DATE				
	WO	2007	0119	39		A 2	A2 20070125			WO 2006-US27855						20060718		
		W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,
			CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
			GE,	GH,	GM,	HN,	HR,	ΗU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KM,	KN,	KP,
			KR,	KZ,	LA,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	LY,	MA,	MD,	MG,	MK,	MN,
		•	MW,	MX,	ΜZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RS,	RU,
			SC,	SD,	SE,	SG,	SK,	SL,	SM,	SY,	ТJ,	TM,	TN,	TR,	TT,	ΤZ,	UA,	UG,
	•		US,	UZ,	VC,	VN,	ZA,	ZM,	ZW									
		RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,
			IS,	IT,	LT,	LU,	LV,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	ВJ,
			CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG,	BW,	GH,
			GM,	KΕ,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	ΤZ,	UG,	ZM,	ZW,	AM,	ΑZ,	BY,
			KG,	ΚZ,	MD,	RU,	TJ,	TM										
	PRIORITY	APP	LN. :	INFO	.:					1	US 20	005-	7006	98P		P 20	00501	718
										_								

US 2005-713907P P 20050901 The present invention features improved processes and organisms for the prodn. of methionine. The invention demonstrates that a .DELTA.metF organism or a .DELTA.metE AmetH organism, for example, mutants of C. glutamicum or E. coli, can use a Me capped sulfide source, e.g., di-Me disulfide (DMDS), as a source of both sulfur and a Me group, bypassing the need for MetH/MetE and MetF activity and the need to reduce sulfate, for the synthesis of methionine. Also described in this patent are data implicating MetY (also called MetZ) as an enzyme that incorporates a Me capped sulfide source, e.g., DMDS, into methionine. A .DELTA.metF .DELTA.metB strain of C. glutamicum can use a Me capped sulfide source, e.g., DMDS, as a source of both sulfide and a Me group. Furthermore, methionine prodn. by engineered prototrophic organisms that

overproduce O-acetyl-homoserine was improved by the addn. of a Me capped

ANSWER 2 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

sulfide source, e.g., DMDS.

ACCESSION NUMBER:

2005:1026863 HCAPLUS

DOCUMENT NUMBER:

143:340637

TITLE:

Lactobacillus acidophilus nucleic acid sequences encoding stress-related proteins and their uses

INVENTOR(S):

Klaenhammer, Todd Robert; Altermann, Eric;

Azcarate-Peril, Andrea; Mcauliffe, Olivia; Russell, W.

Michael

PATENT ASSIGNEE(S):

North Carolina State University, USA

SOURCE:

.PCT Int. Appl., 720 pp.

DOCUMENT TYPE:

Patent

CODEN: PIXXD2

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

DATE PATENT NO. KIND APPLICATION NO. DATE ---**-**----------A2 WO 2005086794 20050922 WO 2005-US7506 WO 2005086794 A3 20061221

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,

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             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM,
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             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
            MR, NE, SN, TD, TG
    US 2005250135
                                20051110
                                            US 2005-74176
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                                                                   20050307
PRIORITY APPLN. INFO.:
                                            US 2004-551161P
                                                                P 20040308
                                            US 2005-74176
                                                                A 20050307
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The current invention provides 185 stress-related nucleic acid mols. and AB their encoded polypeptides and fragments and variants thereof prepd. from Lactobacillus acidophilus. The stress-related proteins include heat and cold shock proteins, acid and alk. tolerance proteins, osmotic and oxidative stress-related proteins, and starvation-induced proteins. In addn., stress-related fusion proteins, antigenic peptides, and anti-stress-related antibodies are encompassed. The invention also provides recombinant expression vectors contg. a nucleic acid mol. of the invention and host cells into which the expression vectors have been introduced. Methods for producing the polypeptides and methods of use for the polypeptides of the invention are further disclosed.

ANSWER 3 OF 9 MEDLINE on STN DUPLICATE 1

ACCESSION NUMBER: 2005287400 MEDLINE DOCUMENT NUMBER: PubMed ID: 15933025

TITLE: Effects of deregulation of methionine biosynthesis on

methionine excretion in Escherichia coli.

Usuda Yoshihiro; Kurahashi Osamu AUTHOR:

Fermentation & Biotechnology Laboratories, Institute of Life Sciences, Ajinomoto Co. Inc., 1-1 Suzuki-cho, CORPORATE SOURCE:

Kawasaki-ku, Kawasaki-shi 210-8681, Japan..

yoshihiro usuda@ajinomoto.com

SOURCE: Applied and environmental microbiology, (2005 Jun) Vol. 71,

No. 6, pp. 3228-34.

Journal code: 7605801. ISSN: 0099-2240.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200508

ENTRY DATE: Entered STN: 4 Jun 2005

> Last Updated on STN: 15 Aug 2005 Entered Medline: 11 Aug 2005

AB Several regulators of methionine biosynthesis have been reported in Escherichia coli, which might represent barriers to the production of excess 1-methionine (Met). In order to examine the effects of these factors on Met biosynthesis and metabolism, deletion mutations of the methionine repressor (metJ) and threonine biosynthetic (thrBC) genes were introduced into the W3110 wild-type strain of E. coli. Mutations of the metK gene encoding S-adenosylmethionine synthetase, which is involved in Met metabolism, were detected in 12 norleucine-resistant mutants. Three of the mutations in the metK structural gene were then introduced into metJ and thrBC double-mutant strains; one of the resultant strains was found to accumulate 0.13 g/liter Met. Mutations of the metA gene encoding homoserine succinyltransferase were detected in alpha-methylmethionine-resistant mutants, and these mutations were found to encode feedback-resistant enzymes in a 14C-labeled homoserine assay. Three metA mutations were introduced, using expression plasmids, into an E. coli strain that was shown to accumulate 0.24 g/liter Met. Combining mutations that affect the deregulation of Met biosynthesis and metabolism is therefore an effective approach for the production of Met-excreting strains.

L2 ANSWER 4 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1996:688432 HCAPLUS

DOCUMENT NUMBER: 126:72244

TITLE: Genetic tools for selective labeling of proteins with

.alpha.-15N-amino acids

AUTHOR(S): Waugh, David S.

CORPORATE SOURCE: Dep. Physical Chem., Roche Research Cent., Nutley, NJ,

07110, USA

SOURCE: Journal of Biomolecular NMR (1996), 8(2), 184-192

CODEN: JBNME9; ISSN: 0925-2738

PUBLISHER: ESCOM
DOCUMENT TYPE: Journal
LANGUAGE: English

AB A collection of genetic tools that can be used to manipulate amino acid metab. in Escherichia coli is described. The set comprises 21 strains of bacteria, each contg. a different genetic defect that is closely linked to a selectable transposon marker. These tools can be used to construct strains of E. coli with ideal genotypes for residue-specific, selective labeling of proteins with nearly any 15N-amino acid. By using strains which have been modified to contain the appropriate genetic lesions to control amino acid biosynthesis, diln. of the isotype by endogenous amino acid biosynthesis and scrambling of the label to other types of residues can be avoided.

L2 ANSWER 5 OF 9 BIOTECHDS COPYRIGHT 2007 THE THOMSON CORP. on STN

ACCESSION NUMBER: 1990-11344 BIOTECHDS

TITLE: S-metabolism of methionine-rich yeasts;

Candida guilliermondii, Candida utilis, Rhodotorula

glutinis, Saccharomyces carlsbergensis single cell protein

AUTHOR: Halasz A; Matrai B; Muayad A

LOCATION: Central Food Research, Herman Otto ut. 15, H-1022 Budapest,

Hungary.

SOURCE: Acta Aliment.Acad.Sci.Hung.; (1989) 18, 4, 361-85

CODEN: AAASCO

DOCUMENT TYPE: Journal LANGUAGE: English

Candida guilliermondii CBS 5256, Candida utilis CBS 5609, Rhodotorula glutinis CBS 315 and Saccharomyces carlsbergensis were subjected to mild mutagenesis using UV-irradiation or nitrite, and mutants were selected on the basis of increased sulfate requirement. About 25% of these mutants showed an increased methionine content. They were more sensitive to norleucine (a methionine antagonist) than wild-type strains, suggesting that their higher Met content was not due to homoserine-succinyltransferase (EC-2.3.1.46) derepression. The mutants responded to a higher methyl donor concentration by enhanced growth. The lipoic acid concentration of the yeast increased in parallel to the augmentation of sulfur-containing amino acids. The concentration of these amino acids was sensitive to aeration intensity, and dropped at levels above 200 mmol O2/hr.l. This was explained by the utilization of the sulfur derived from sulfate reduction being converted to sulfide by yeast sulfate-reductase, of which lipoic acid is a prosthetic group. Sulfate was a better S-source than methionine. (19 ref)

L2 ANSWER 6 OF 9 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights

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ACCESSION NUMBER: 78044402 EMBASE

DOCUMENT NUMBER: 1978044402

TITLE: Repression of the tyrosine, lysine, and methionine

biosynthetic pathways in a hisT mutant of

Salmonella typhimurium.

AUTHOR: Brown B.A.; Lax S.R.; Liang L.; et al.

CORPORATE SOURCE: Clayton Found. Biochem. Inst., Univ. Texas, Austin, Tex.

78712, United States

SOURCE: Journal of Bacteriology, (1977) Vol. 129, No. 2, pp.

1168-1170.

CODEN: JOBAAY

DOCUMENT TYPE:

Journal

FILE SEGMENT:

004 Microbiology

022 Human Genetics

LANGUAGE:

English

ANSWER 7 OF 9 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights

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ACCESSION NUMBER:

78178718 EMBASE

DOCUMENT NUMBER:

1978178718

TITLE:

Norleucine accumulation by a norleucine-resistant

mutant of Serratia marcescens.

AUTHOR:

Kisumi M.; Sugiura M.; Chibata I.

CORPORATE SOURCE:

Res. Lab. Appl. Biochem., Tanabe Seiyaku Co. Ltd, Osaka,

SOURCE:

Applied and Environmental Microbiology, (1977) Vol. 34, No.

2, pp. 135-138. .

CODEN: AEMIDF

COUNTRY:

United States

DOCUMENT TYPE:

Journal

FILE SEGMENT:

004 Microbiology

LANGUAGE:

English

L2 ANSWER 8 OF 9 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights

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ACCESSION NUMBER:

78036460 EMBASE

DOCUMENT NUMBER:

1978036460

TITLE:

Sulfur amino acid auxotrophy in Micrococcus species

isolated from human skin.

AUTHOR:

Farrior J.W.; Kloos W.E.

CORPORATE SOURCE:

Dept. Genet., North Carolina State Univ., Raleigh N.C.

27607, United States

SOURCE:

Canadian Journal of Microbiology, (1976) Vol. 22, No. 12,

pp. 1680-1690. .

CODEN: CJMIAZ

DOCUMENT TYPE:

Journal

FILE SEGMENT:

004 Microbiology

LANGUAGE: English

Since methionine and (or) cysteine are required by a large percentage of natural auxotrophic Micrococcus strains isolated from human skin, investigations were directed to determine the specific enzymes affected in sulfur amino acid biosynthesis. Known intermediates in the interrelated cysteine methionine biosynthetic pathways were tested as growth stimulants. Based on these growth studies, sulfur amino acid auxotrophs were grouped into three cysteine classes and five methionine classes. Selected auxotrophs of M. luteus had deficiencies in ATP sulfurylase (EC 2.7.7.4) and adenosine 5 sulfatophosphate (APS) kinase (EC 2.7.1.25), sulfite reductase (EC 1.8.1.2), serine transacetylase (EC 2.3.1.30), or .beta. cystathionase (EC 4.4.1.8) activity; auxotrophs of M. lylae had deficiencies in sulfite reductase and serine transacetylase, .beta. cystathionase, or N5,N10 methyltetrahydrofolate reductase (EC 1.1.1.68) activity; all auxotrophs of M. sedentarius tested had deficiencies in N5,N10 methyltetrahydrofolate reductase activity; auxotrophs of M. nishinomiyaensis had deficiencies in adenosine 3 phosphate 5 sulfatophosphate (PAPS) reductase, sulfite reductase, serine transacetylase, or N5,N10 methyltetrahydrofolate reductase activity; auxotrophs of M. varians had deficiencies in APS kinase, PAPS reductase, sulfite reductase, homoserine O transsuccinylase, .beta. cystathionase, or N5,N10 methyltetrahydrofolate reductase activity; auxotrophs of M. kristinae had deficiencies in serine transacetylase or cystathionine .gamma. synthase (EC 4.2.99.9) activity; auxotrophs of M. roseus had deficiencies in PAPS reductase, sulfite reductase, or serine transacetylase activity. Results of studies with various mutagens

suggested that sulfur amino acid auxotrophy was primarily the result of a single base substitution in usually one or two of the genes controlling biosynthesis. A preliminary study of the amino acid composition of sweat suggested that this important source of nutrients does not contain adequate amounts of cysteine for the growth of cysteine auxotrophs but contains methionine that may be utilized in place of cysteine.

ANSWER 9 OF 9 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights L2

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ACCESSION NUMBER: 74118263 EMBASE

DOCUMENT NUMBER: 1974118263

TITLE: The control of homoserine O transsuccinylase in a

methionine requiring mutant of the blue green

alga Anacystis nidulans.

Delaney S.F.; Dickson A.; Carr N.G. **AUTHOR:**

Dept. Biochem., Univ. Liverpool, United Kingdom CORPORATE SOURCE:

Journal of General Microbiology, (1973) Vol. 79, No. 1, pp. SOURCE:

89-94.

CODEN: JGMIAN

DOCUMENT TYPE: Journal

FILE SEGMENT: 022 Human Genetics

> 029 Clinical Biochemistry

LANGUAGE: English

=> s MetA and (101 or 294)

604 META AND (101 OR 294)

=> s 16 and (variant? or mutant?)

27 L6 AND (VARIANT? OR MUTANT?)

=> s 17 and escherichia

1 L7 AND ESCHERICHIA

=> d 18 ibib ab

ANSWER 1 OF 1 BIOTECHDS COPYRIGHT 2007 THE THOMSON CORP. on STN

ACCESSION NUMBER: 2004-15360 BIOTECHDS

TITLE: New mutants of homoserine-transsuccinylase, useful,

when expressed in microorganisms, for production of

methionine and S-adenosyl-methionine, also related nucleic

recombinant enzyme production via plasmid expression in host cell for use in feed-additive and disease therapy

AUTHOR: WINTERHALTER C; LEONHARTSBERGER S; PFEIFFER K; BAUER B

PATENT ASSIGNEE: CONSORTIUM ELEKTROCHEM IND GMBH

PATENT INFO: DE 10247437 29 Apr 2004 APPLICATION INFO: DE 2002-1047437 11 Oct 2002

PRIORITY INFO: DE 2002-1047437 11 Oct 2002; DE 2002-1047437 11 Oct 2002

DOCUMENT TYPE: Patent LANGUAGE: German

OTHER SOURCE: WPI: 2004-348936 [33]

DERWENT ABSTRACT:

NOVELTY - Homoserine-transsuccinylase (MetA) that contains a mutation, relative to the wild type, and has reduced sensitivity to L-methionine and S-adenosyl-methionine (SAM) is new.

DETAILED DESCRIPTION - Homoserine-transsuccinylase (MetA) that contains a mutation, relative to the wild type, and has reduced sensitivity to L-methionine and S-adenosyl-methionine (SAM) is new. The mutation is: (a) of Asp in sequence (1), present between positions 90 and 115; or (b) of Tyr in sequence (2), present between positions 285 and 310. Where Asp-Gly-X-X-Thr-Gly-Ala-Pro (1) Tyr-Gln-X-Thr-Pro (2). X =any amino acid. INDEPENDENT CLAIMS are also included for: (1) the metA alleles (I) that encode the new mutants; (2) plasmids that contain (I); (3) microorganisms that include a

feedback-resistant (I); and (4) method for preparing L-Met or SAM by culturing the organisms of (3).

BIOTECHNOLOGY - Preferred Enzymes: The new metA mutants have at least double, particularly 50 times, the resistance (expressed as Ki) of the wild-type enzyme with respect to Met and SAM. The specification includes a table listing suitable mutants; e.g. (a) GAC (Asp) as codon 101 but TAC (Tyr) at codon 294 is replaced by TGC (Cys); CTC (Leu); GCC (Ala) or is absent; and (b) TAC as codon 294 but codon 101 is AAC (Asn), CAC (His), TGT (Cys) or AGC (Ser). Preferred Organism: This is a Gram-negative bacterium, specifically Escherichia coli. Preparation: The new metA alleles are produced by standard methods of random or targeted mutagenesis, using the wild-type sequence as template.

ACTIVITY - Antidepressant; Hepatotropic; Antiarthritic. No biological data is given.

MECHANISM OF ACTION - None given.

USE - The mutant MetA is expressed in host cells, for production of L-Met (used as feed additive) and SAM (used to treat depression, liver disease and arthritis) (claimed); also for production of Met-containing peptides and metabolites of Met and SAM such as polyamines, lipoic acid; biotin and quinones.

ADVANTAGE - The new mutants are less sensitive than the wild type to feedback inhibition by Met and SAM, so provide improved yields of these compounds. The wild-type MetA from Escherichia coli W3110 retained 2% of its activity in presence of 1 mM Met, and had Ki 0.05 mM; compare 96% and 11 mM for the mutant with Cys instead of Tyr at position 294. For 1 mM SAM corresponding figures were 0.5% and 0.2 mM for the wild type and 92% and 10 mM for the mutant.

EXAMPLE - Plasmid pKP413GAP contains the metA (homoserine-transsuccinylase) gene of Escherichia coli W3110 under control of the GAPDH promoter. It was subjected to site-specific mutation to convert codon 294 (TAC (Tyr) to TGC (Leu), primer sequences reproduced. The resulting 4.3 kb fragment was used to transform E. coli DH5alpha, for production of the mutant enzyme. This was resistant to inhibition by both methionine and its S-adenosyl derivative. (22 pages)

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FILE 'SCISEARCH' ENTERED AT 10:34:56 ON 20 FEB 2007 Copyright (c) 2007 The Thomson Corporation

=> s homoserine transsuccinylase (mutant? or variant? or mutation?)
MISSING OPERATOR 'CCINYLASE (MUTANT?'
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nested terms that are not separated by a logical operator.

=> dup rem l1

PROCESSING COMPLETED FOR L1

L2 15 DUP REM L1 (12 DUPLICATES REMOVED)

=> s 12 and Escherichia

L3 9 L2 AND ESCHERICHIA

=> d 13 1-9 ibib ab

L3 ANSWER 1 OF 9 MEDLINE ON STN ACCESSION NUMBER: 95173116 MEDLINE DOCUMENT NUMBER: PubMed ID: 7868613

TITLE: Heat shock-dependent transcriptional activation of the metA

gene of Escherichia coli.

AUTHOR: Biran D; Brot N; Weissbach H; Ron E Z

CORPORATE SOURCE: Department of Molecular Microbiology and Biotechnology,

Tel-Aviv University, Israel.

SOURCE: Journal of bacteriology, (1995 Mar) Vol. 177, No. 5, pp.

1374-9.

Journal code: 2985120R. ISSN: 0021-9193.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199503

ENTRY DATE: Entered STN: 7 Apr 1995

Last Updated on STN: 6 Feb 1998 Entered Medline: 27 Mar 1995

AB In Escherichia coli, the growth rate at elevated temperatures is controlled by the availability of endogenous methionine, which is limited because of the temperature sensitivity of the metA gene product, homoserine transsuccinylase (HTS). In order to

determine the relationship between this control mechanism and the heat shock response, we estimated the cellular levels of HTS during heat shock by Western (immunoblot) analysis and found an increase following induction by temperature shift and by addition of ethanol or cadmium ions. The elevated level of HTS was a result of transcriptional activation of the metA gene. This activation was heat shock dependent, as it did not take place in rpoH mutants, and probably specific to the metA gene, as another gene of the methionine regulon (metE) was not activated. These results suggest a metabolic link between the two systems that control the response of E. coli to elevated temperatures: the metA gene, which codes for the enzyme responsible for regulating cell growth as a function of temperature elevation (HTS), is transcriptionally activated by the heat shock response.

L3 ANSWER 2 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2007:172138 HCAPLUS

TITLE: Identification of Catalytic Cysteine, Histidine, and

Lysine Residues in Escherichia coli

Homoserine Transsuccinylase

AUTHOR(S): Ziegler, Katharine; Noble, Schroeder M.; Mutumanje,

Elissa; Bishop, Barney; Huddler, Donald P.; Born,

Timothy L.

CORPORATE SOURCE: Department of Chemistry Biochemistry, George Mason

University, Manassas, VA, 20110, USA

SOURCE: Biochemistry ACS ASAP

CODEN: BICHAW; ISSN: 0006-2960

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal LANGUAGE: English

AB Homoserine transsuccinylase catalyzes the

succinylation of homoserine in several bacterial species, the first unique step in methionine biosynthesis in these organisms. The enzyme from Escherichia coli is reported to be a dimer and uses a ping-pong catalytic mechanism involving transfer of succinate from succinyl-CoA to an enzyme nucleophile, followed by transfer to homoserine to form O-succinylhomoserine. Site-directed mutagenesis and steady-state kinetics were used to identify three amino acids that participate in catalysis. Mutation of cysteine-142 to serine or alanine eliminated all measurable activity, suggesting this amino acid acts as the catalytic nucleophile. Cysteine nucleophiles are often deprotonated by histidine residues, and histidine-235 was identified as the sole absolutely conserved histidine residue among family members. This residue was mutated to both alanine and asparagine, and no activity was obsd. with either mutant. Lysine-47 had been previously identified as an essential residue. Mutation of this amino acid to arginine reduced catalytic activity by greater than 90%, while mutation to alanine yielded an enzyme with <1% of wild-type activity. A pH-rate profile of the K47R mutant demonstrated that this amino acid participates in the first half reaction. The data presented here provide the first detailed description of the homoserine transsuccinylase active site and provide a framework for addnl. mechanistic characterization of this enzyme.

L3 ANSWER 3 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:1239041 HCAPLUS

DOCUMENT NUMBER: 144:2275

TITLE: Construction of microorganism containing recombinant

homoserine transsuccinylase with

altered feedback sensitivity and recombinant

S-adenosylmethionine synthetase with reduced activity

for the production of methionine

INVENTOR(S): Bestel-Corre, Gwenaeelle Anne Lise; Chateau, Michel;

Figge, Rainer Martin; Raynaud, Celine; Soucaille,

Philippe Noeel Paul

PATENT ASSIGNEE(S): Metabolic Explorer, Fr.

SOURCE: PCT Int. Appl., 72 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:
FAMILY ACC. NUM. COUNT:

Engl

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE _ _ _ _ ----------WO 2004-IB1901 WO 2005111202 A1 20051124 20040512 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG WO 2005108561 **A2** 20051117 WO 2005-EP52180 20050512 WO 2005108561 A3 20060720 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZM ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG EP 1747269 A2 20070131 EP 2005-742717 20050512 AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, LV, MK, YU PRIORITY APPLN. INFO.: WO 2004-IB1901 A 20040512 WO 2005-EP52180 W 20050512 OTHER SOURCE(S): CASREACT 144:2275; MARPAT 144:2275 The present invention relates to the use of recombinant homoserine transsuccinylase with altered sensitivity to feedback inhibitors S-adenosylmethionine and methionine (MetA*) and optionally, recombinant S-adenosylmethionine synthetase with reduced activity (MetK*) for the prodn. of methionine, its precursors or derivs. thereof. More specifically, the authors isolated Escherichia coli mutants contg. homoserine transsuccinylase which show decreased feedback-sensitivity towards S-adenosylmethionine and methionine. E. coli mutants contg. S-adenosylmethionine synthetase with reduced activity were also isolated. Construction of E. coli strains for the prodn. of O-succinylhomoserine or methionine by combining feed-back resistant MetA alleles with MetK alleles with decreased activity is described. Fermn. of E. coli prodn. strains and

L3 ANSWER 4 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

6

anal. of yield is reported.

ACCESSION NUMBER:

REFERENCE COUNT:

2005:1220814 HCAPLUS

DOCUMENT NUMBER:

143:474228

TITLE:

Construction of microbial recombinant homoserine transsuccinylase with

altered feedback sensitivity and S-adenosyl methionine

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

synthetase with reduced activity for the production of

methionine

INVENTOR(S): Bestel-Corre, Gwenaeelle; Chateau, Michel; Figge,

Rainer Martin; Raynaud, Celine; Soucaille, Philippe

Noel Paul

PATENT ASSIGNEE(S): Metabolic Explorer, Fr. SOURCE: PCT Int. Appl., 46 pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

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APPLICATION NO.
         PATENT NO.
                                             KIND
                                                           DATE
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         WO 2005108561
                                                                                WO 2005-EP52180
                                               A2
                                                           20051117
                                                                                                                           20050512
         WO 2005108561
                                               Α3
                                                           20060720
               W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
                RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
               2005111202
A1 20051124 WO 2004-IB1901 20040512
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
         WO 2005111202
                                               A1
                                                           20051124
                                                                              WO 2004-IB1901
                                                          20070131
         EP 1747269
                                                                              EP 2005-742717
                                               A2
                                                                                                                           20050512
                       AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA,
                       HR, LV, MK, YU
PRIORITY APPLN. INFO.:
                                                                                WO 2004-IB1901
                                                                                                                     A 20040512
                                                                                WO 2005-EP52180
                                                                                                                     W 20050512
OTHER SOURCE(S):
                                             CASREACT 143:474228; MARPAT 143:474228
         The present invention relates to the use of recombinant homoserine
         transsuccinylase with altered feedback sensitivity (MetA*) and
         eventually, recombinant S-adenosyl methionine synthetase with reduced
         activity (MetK*) for the prodn. of methionine, its precursors or derivs.
         thereof. More specifically, Escherichia coli mutants
         contg. homoserine transsuccinylase with decreased
         feedback sensitivity towards methionine and S-adenosylmethionine were
         isolated. E. coli mutants contg. S-adenosyl methionine
         synthetase with reduced activity were also isolated. Construction of E.
        coli strains for the prodn. of O-succinylhomoserine or methionine by
        combined feed-back resistant MetA alleles with MetK alleles with decreased
        activity is described. Fermn. of E. coli prodn. strains and anal. of
        yield is reported.
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L3 ANSWER 5 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

140:387796

ACCESSION NUMBER: 2004:371078 HCAPLUS

DOCUMENT NUMBER:

TITLE: Methionine and SAM feedback-resistant homoserine

transsuccinylases with modified C-terminus Leonhartsberger, Susanne; Pfeiffer, Kerstin;

Winterhalter, Christoph; Bauer, Brigitte

PATENT ASSIGNEE(S): Consortium fuer Elektrochemische Industrie G.m.b.H.,

Germany

SOURCE: PCT Int. Appl., 33 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

INVENTOR (S):

PAT	PATENT NO.				KIND DATE			APPLICATION NO.						DATE			
		- -		•				-									
WO	2004	0380	13		A2	2004	0506	W	0 2	003-	EP11	486		2	0031	016	
WO	2004	0380	13		· A3	2004	0624										
	W:	CA,	CN,	JP,	RU,	US											
	RW:	ΑT,	BE,	BG,	CH,	CY, CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	
		IT,	LU,	MC,	NL,	PT, RO,	SE,	SI,	SK,	TR							
DE	1024	9642			A1	2004	0513	D	E 2	002-	1024	9642		2	0021	024	
EP	1570	066			A2	2005	0907	E	P 2	003-	7694	05		2	0031	016	
EP	1570	066			B1	2006	1227										
	R:	ΑT,	BE,	CH,	DE,	DK, ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
		ΙE,	SI,	FI,	RO,	CY, TR,	BG,	CZ,	EE,	HU,	SK						
CN	1705	751			A	2005	1207	C	N 2	003-	8010	1894		2	0031	016	
JP	2006	50356	58		T	2006	0202	J	P 2	004-	5458	67		2	0031	016	
AT	34954	46			T	2007	0115	Α	т 2	003-	7694	05		2	0031	016	
· US	2006	1601	73		A1	2006	0720	U	S 2	005-	5308	44		2	0050	408	
PRIORITY	Y APP	LN.	INFO.	. :				D	E 2	002-	1024	9642		A 2	0021	024	
								W	0 2	003-	EP11	486	1	W 2	0031	016	

AB The invention relates to a homoserine transsuccinylase
, which exhibits reduced sensitivity towards L-methionine or SAM in
comparison with a homoserine transsuccinylase
wild-type enzyme, whereby the latter comprises an amino acid sequence
contg. a TyrGlnXaaThrPro sub-sequence, the Thr of said sub-sequence lying
between positions 285 and 310 of the amino acid sequence. The inventive
homoserine transsuccinylase is characterized in that in
comparison with the wild-type enzyme at least 2 amino acids are modified,
said modification taking place in the Thr of the sub-sequence or in the
C-terminal. Thus, exts. of E. coli contg. metA gene mutants
were analyzed for homoserine transsuccinylase activity
in the presence of 1 mM Met or SAM. The wild-type enzyme retains 2% and
0.5% activity, resp. One of the mutants exhibited 95% activity
under these circumstances. The Ki for Met was 16 mM and for SAM 9 mM.

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L3 ANSWER 6 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN
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ACCESSION NUMBER: 2004:349595 HCAPLUS

DOCUMENT NUMBER: 140:370810

TITLE: Feedback-resistant homoserine

transsuccinylase mutants,

microorganisms producing them, and their use in

production of methionine and SAM

INVENTOR(S): Winterhalter, Christoph; Leonhartsberger, Susanne;

Pfeiffer, Kerstin; Bauer, Brigitte

PATENT ASSIGNEE(S): Consortium fuer Elektrochemische Industrie G.m.b.H.,

Germany

SOURCE: Ger. Offen., 22 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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20040429 DE 2002-10247437
    DE 10247437
                        A1
                                                                  20021011
    WO 2004035617
WO 2004035617
                               20040429 WO 2003-EP10978
                        A2
                                                                  20031002
                        A3
                               20040617
        W: CA, CN, JP, RU, US
        RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
            IT, LU, MC, NL, PT, RO, SE, SI, SK, TR
                         A2 20050706 EP 2003-767502
    EP 1549754
                                                                  20031002
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, SK
    CN 1703517
                             20051130
                                          CN 2003-80101208
                                                                  20031002
                        Α
    JP 2006516092
                         Т
                               20060622
                                           JP 2004-544072
                                                                  20031002
                                           DE 2002-10247437 A 20021011
WO 2003-EP10978 W 20031002
PRIORITY APPLN. INFO.:
```

Homoserine transsuccinylase, which contains at least AB one mutation compared to a homoserine transsuccinylase wild type enzyme and compared to the wild type enzyme shows a reduced sensitivity to L-methionine or SAM is disclosed. The wild-type enzyme contains a partial sequence AspGlyXaaXaaXhrGlyAlaPro between residues 90 and 115 and a partial sequence TyrGlnXaaThrPro between residues 285 and 310. The mutations comprise an amino acid exchange of Asp in AspGlyXaaXaaXaaThrGlyAlaPro or an amino acid exchange of Tyr in TyrGlnXaaThrPro. Thus, the Y294C mutant of E. coli MetA exhibits 96% activity in the presence of 1 mM Met while the wild-type enzyme is almost totally inhibited. The Ki for Met in the mutant is 11 mM, for Met in the wild-type, 0.05 mM. The same mutant show 92% activity in the presence of 1 mM SAM and a Ki of 10 mM, while the wild-type enzyme shows negligible activity and Ki of 0.2 mM.

ANSWER 7 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2000:342198 HCAPLUS

DOCUMENT NUMBER:

133:3756

TITLE:

L-methionine and its preparation with transgenic

Escherichia coli mutants with

defective repressor and enhanced homoserine

transsuccinylase activity

INVENTOR(S):

Usuta, Yoshihiro; Kurahashi, Osamu

PATENT ASSIGNEE(S):

Ajinomoto Co., Inc., Japan

SOURCE:

Jpn. Kokai Tokkyo Koho, 23 pp.

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

L3

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE						
	JP 2000139471	 А	20000523	JP 1998-326717	19981117						
PRIO	RITY APPLN. INFO.:			JP 1998-326717	19981117						
AB	Described is a meth	od of m	anufg. L-met	hionine by cultivating	a						
	Escherichia coli mu										
	metJ), enhanced hom	oserine	transsuccin	ylase (gene							
	metA) activity, and	, optio	nally, decre	ased S-adenosyl methior	nine						
				mutants may also have	the						
•				gammasynthase and							
•	aspartokinase-homos	erine d	ehydrogenase	II. Also claimed are	the						
	S-adenosyl methioni	ne synt	hetase (metK) mutants with							
	substitution mutati	ons at	27-Arg.fwdar	w.Cys, 296-Ile.fwdarw.S	Ser, .						
	298-Pro.fwdarw.Leu, or a combination of them. The mutants are										
	free of the synergi	stic in	hibition by	L-methionine and S-ader	osyl						
				th improved efficiency	by using the						
	Escherichia coli mu	tants w	as demonstra	ted.	-						

ACCESSION NUMBER: 1984:524135 HCAPLUS

DOCUMENT NUMBER: 101:124135

TITLE: Expression of the metA gene of Escherichia

> coli K-12 in recombinant plasmids Michaeli, Shulamit; Ron, Eliora Z.

CORPORATE SOURCE: George S. Wise Fac. Life Sci., Tel Aviv Univ., Tel

Aviv-Jaffa, Israel

SOURCE: FEMS Microbiology Letters (1984), 23(2-3), 125-9

CODEN: FMLED7; ISSN: 0378-1097

DOCUMENT TYPE: Journal LANGUAGE: English

AUTHOR (S):

The expression of the metA gene for homoserine transsuccinylase [9030-70-0] was studied in wild-type and in deregulated strains of E. coli K-12 carrying the gene on multicopy plasmids. The mol. wt. of the product synthesized by the metA gene was 40,000; the whole enzyme consisted of 2 subunits. In deregulated strains (i.e., those carrying a metJ mutation), the activity of the metA gene was increased 2-fold. Thus, even when metA is cloned onto a multicopy plasmid, it is under the neg. control of the regulatory metJ gene.

ANSWER 9 OF 9 HCAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1982:469190 HCAPLUS

DOCUMENT NUMBER: 97:69190

TITLE: Mechanisms involved in the increased sensitivity of

Escherichia coli to microcin 15m at

42.degree.C

AUTHOR (S): Aguilar, Alfredo; Perez-Diaz, Jose C.; Asensio, Carlos

Inst. Enzimol. Patol. Mol., Madrid, Spain CORPORATE SOURCE: Current Microbiology (1982), 7(2), 83-6 CODEN: CUMIDD; ISSN: 0343-8651 SOURCE:

DOCUMENT TYPE: Journal LANGUAGE: English

E. coli Cells show a markedly increased sensitivity to the antibiotic microcin 15m when briefly treated at 42.degree. as compared to the effect at 37.degree.. Furthermore, mutants resistant to the microcin at 37.degree. become sensitive at 42.degree. at microcin concns. that are inactive at 37.degree.. This effect can be overcome by L-methionine. The mechanism involved seems to be based on an apparent inactivation of the homoserine-O-transsuccinylase activity. As previously established, this enzyme suffers a reversible partial inactivation when the cells are shifted to 42.degree. and the action of the microcin at this temp. seems to bring this process to a virtually irreversible stage. In mixed cultures of the microcin-producing strain and 1 E. coli stain sensitive to the antibiotic, a much stronger growth inhibition of the latter strain was obsd. at 42.degree. than at 37.degree..

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(FILE 'HOME' ENTERED AT 10:34:17 ON 20 FEB 2007)

FILE 'MEDLINE, HCAPLUS, BIOSIS, BIOTECHDS, EMBASE, SCISEARCH' ENTERED AT 10:34:56 ON 20 FEB 2007

L127 S HOMOSERINE TRANSSUCCINYLASE AND (MUTANT? OR VARIANT? OR MUTAT

L2 15 DUP REM L1 (12 DUPLICATES REMOVED)

L3 9 S L2 AND ESCHERICHIA

=> s 13 and codon 101 or 294

32069 L3 AND CODON 101 OR 294

=> s 13 and (codon 101 or codon 294)

0 L3 AND (CODON 101 OR CODON 294)

=> s 13 and (101 or 294)

=> s 13 and Y294C

L7 1 L3 AND Y294C

=> d his

(FILE 'HOME' ENTERED AT 10:34:17 ON 20 FEB 2007)

FILE 'MEDLINE, HCAPLUS, BIOSIS, BIOTECHDS, EMBASE, SCISEARCH' ENTERED AT 10:34:56 ON 20 FEB 2007

L1 27 S HOMOSERINE TRANSSUCCINYLASE AND (MUTANT? OR VARIANT? OR MUTAT

L2 15 DUP REM L1 (12 DUPLICATES REMOVED)

L3 9 S L2 AND ESCHERICHIA

L4 32069 S L3 AND CODON 101 OR 294

L5 0 S L3 AND (CODON 101 OR CODON 294)

L6 0 S L3 AND (101 OR 294)

L7 1 S L3 AND Y294C

=> log y

COST IN U.S. DOLLARS

SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION

CA SUBSCRIBER PRICE -6.24 -6.24

STN INTERNATIONAL LOGOFF AT 10:40:36 ON 20 FEB 2007